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## **CLAIMS**

- 1. A method of diagnosing PRC or a predisposition to developing PRC in a subject, comprising determining a level of expression of EphA4 in a patient derived biological sample, wherein an increase of said level compared to a normal control level of said gene indicates that said subject suffers from or is at risk of developing PRC.
- 2. The method of claim 1, wherein said increase is at least 10% greater than said normal control level.
- 3. The method of claim 1, wherein the expression level is determined by any one method select from group consisting of:
  - (a) detecting the mRNA of EphA4,
  - (b) detecting the protein encoded by EphA4, and
  - (c) detecting the biological activity of the protein encoded by EphA4.
- 4. The method of claim 1, wherein said level of expression is determined by detecting hybridization of EphA4 probe to a gene transcript of said patient-derived biological sample.
  - 5. The method of claim 4, wherein said hybridization step is carried out on a DNA array.
  - 6. The method of claim 1, wherein said biological sample comprises an epithelial cell.
- 7. The method of claim 1, wherein said biological sample comprises PRC cell.
  - 8. The method of claim 7, wherein said biological sample comprises an epithelial cell from a PRC.
  - 9. A method of screening for a compound for treating or preventing PRC, said method comprising the steps of:
  - a) contacting a test compound with a polypeptide encoded by EphA4;
    - b) detecting the binding activity between the polypeptide and the test compound;

and

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- c) selecting a compound that binds to the polypeptide.
- 10. A method of screening for a compound for treating or preventing PRC, said method comprising the steps of:
  - a) contacting a candidate compound with a cell expressing EphA4; and
  - b) selecting a compound that reduces the expression level of EphA4.
- 11. The method of claim 10, wherein said cell comprises a prostate cancer cell.
- 12. A method of screening for a compound for treating or preventing PRC, said method comprising the steps of:
  - a) contacting a test compound with a polypeptide encoded by EphA4;
  - b) detecting the biological activity of the polypeptide of step (a); and
  - c) selecting a compound that suppresses the biological activity of the polypeptide in comparison with the biological activity detected in the absence of the test compound.
- 15 13. The method of claim12, wherein the biological activity is tyrosine kinase activity.
  - 14. A method of screening for compound for treating or preventing PRC, said method comprising the steps of:
    - a) contacting a test compound with a cell into which a vector comprising the transcriptional regulatory region of EphA4 genes and a reporter gene that is expressed under the control of the transcriptional regulatory region has been introduced,
    - b) measuring the expression or activity of said reporter gene; and
    - c) selecting a compound that reduces the expression or activity level of said reporter gene, as compared to a level in the absence of the test compound.
- 25 15. A method of treating or preventing PRC in a subject comprising administering to said subject an antisense composition, said composition comprising a nucleotide sequence complementary to a coding sequence of EphA4.
  - 16. A method of treating or preventing PRC in a subject comprising administering to

- said subject a siRNA composition, wherein said composition reduces the expression of EphA4.
- 17. The method of claim 16, wherein said siRNA comprises a sense nucleic acid and an anti-sense nucleic acid of *EphA4*.
- 5 18. The method of claim 17, wherein the siRNA comprises a ribonucleotide sequence corresponding to a sequence consisting of SEQ ID NO: 10 as the target sequence.
  - 19. The method of claim 18, said siRNA has the general formula 5'-[A]-[B]-[A']-3', wherein [A] is a ribonucleotide sequence corresponding to a sequence consisting of nucleotides of SEQ ID NO: 10.
- [B] is a ribonucleotide sequence consisting of about 3 to about 23 nucleotides, and [A'] is a ribonucleotide sequence consisting of the complementary sequence of [A].
  - 20. The method of claim 16, wherein said composition comprises a transfection-enhancing agent.

- A method of treating or preventing PRC in a subject comprising the step of
  administering to said subject a pharmaceutically effective amount of an antibody or
  fragment thereof that binds to a protein encoded by EphA4.
  - 22. A method of treating or preventing PRC in a subject comprising administering to said subject a vaccine comprising a polypeptide encoded by EphA4 or an immunologically active fragment of said polypeptide, or a polynucleotide encoding the polypeptide.
  - 23. A method of treating or preventing PRC in a subject, said method comprising the step of administering a compound that is obtained by the method according to any one of claims 9-14.
- 24. A composition for treating or preventing PRC, said composition comprising a

  pharmaceutically effective amount of an antisense polynucleotide or siRNA against
  a EphA4 as an active ingredient, and a pharmaceutically acceptable carrier.

- 25. The composition of claim 24, wherein said siRNA comprises the nucleotide sequence consisting of SEQ ID NO: 10 as the target sequence.
- 26. A composition for treating or preventing PRC, said composition comprising a pharmaceutically effective amount of an antibody or fragment thereof that binds to a protein encoded by EphA4 as an active ingredient, and a pharmaceutically acceptable carrier.
- 27. A composition for treating or preventing PRC, said composition comprising a pharmaceutically effective amount of the compound selected by the method of any one of claims 9-14 as an active ingredient, and a pharmaceutically acceptable carrier.
- 28. A method for treating or preventing pancreatic cancer in a subject comprising administering to said subject a composition comprising a siRNA of *EphA4*.

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- 29. The method of claim 28, wherein said siRNA comprises a sense nucleic acid and an anti-sense nucleic acid of *EphA4*.
- 30. The method of claim 28, wherein the pancreatic cancer is an pancreatic ductal adenocarcinoma (PDACa).
  - 31. The method of claim 29, wherein the siRNA comprises a ribonucleotide sequence corresponding to a sequence consisting of SEQ ID NO: 10 as the target sequence.
  - 32. The method of claim 31, said siRNA has the general formula 5'-[A]-[B]-[A']-3', wherein [A] is a ribonucleotide sequence corresponding to a sequence consisting of nucleotides of SEQ ID NO: 10.
    - [B] is a ribonucleotide sequence consisting of about 3 to about 23 nucleotides, and [A'] is a ribonucleotide sequence consisting of the complementary sequence consisting of [A].
  - 33. The method of claim 28, wherein said composition comprises a transfection-enhancing agent.
  - 34. A double-stranded molecule comprising a sense strand and an antisense strand,

wherein the sense strand comprises a ribonucleotide sequence corresponding to a target sequence consisting of SEQ ID NO: 10, and wherein the antisense strand comprises a ribonucleotide sequence which is complementary to said sense strand, wherein said sense strand and said antisense strand hybridize to each other to form said double-stranded molecule, and wherein said double-stranded molecule, when introduced into a cell expressing the *EphA4* gene, inhibits expression of said gene.

- 35. The double-stranded molecule of claim 34, wherein said target sequence comprises at least about 10 contiguous nucleotides from the nucleotide sequence consisting of SEQ ID NO: 1.
- 10 36. The double-stranded molecule of claim 35, wherein said target sequence comprises from about 19 to about 25 contiguous nucleotides from the nucleotide sequence consisting of SEQ ID NO: 1.

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- 37. The double-stranded molecule of claim 36, wherein said double-stranded molecule is a single ribonucleotide transcript comprising the sense strand and the antisense strand linked via a single-stranded ribonucleotide sequence.
- 38. The double-stranded molecule of claim 35, wherein the double-stranded molecule is an oligonucleotide of less than about 100 nucleotides in length.
- 39. The double-stranded molecule of claim 38, wherein the double-stranded molecule is an oligonucleotide of less than about 75 nucleotides in length.
- 20 40. The double-stranded molecule of claim 39, wherein the double-stranded molecule is an oligonucleotide of less than about 50 nucleotides in length.
  - 41. The double-stranded molecule of claim 40, wherein the double-stranded molecule is an oligonucleotide of less than about 25 nucleotides in length.
- 42. The double-stranded polynucleotide of claim 41, wherein the double stranded molecule is an oligonucleotide of between about 19 and about 25 nucleotides in length.

- 43. A vector encoding the double-stranded molecule of claim 35.
- 44. The vector of claim 43, wherein the vector encodes a transcript having a secondary structure and comprises the sense strand and the antisense strand.
- 45. The vector of claim 44, wherein the transcript further comprises a single-stranded ribonucleotide sequence linking said sense strand and said antisense strand.
- 46. A vector comprising a polynucleotide comprising a combination of a sense strand nucleic acid and an antisense strand nucleic acid, wherein said sense strand nucleic acid comprises nucleotide sequence consisting of SEQ ID NO: 10, and said antisense strand nucleic acid consists of a sequence complementary to the sense strand.
- 10 47. The vector of claim 46, wherein said polynucleotide has the general formula 5'-[A]-[B]-[A']-3'

wherein [A] is a nucleotide sequence consisting of SEQ ID NO: 10; [B] is a nucleotide sequence consisting of about 3 to about 23 nucleotides; and [A'] is a nucleotide sequence complementary to [A].

- A pharmaceutical composition for treating or preventing pancreatic cancer comprising a pharmaceutically effective amount of a small interfering RNA (siRNA) of *EphA4* as an active ingredient, and a pharmaceutically acceptable carrier.
  - 49. The pharmaceutical composition of claim 48, wherein the siRNA comprises a nucleotide sequence consisting of SEQ ID NO: 10 as the target sequence.
- 50. The composition of claim 49, wherein the siRNA has the general formula 5'-[A]-[B]-[A']-3' wherein [A] is a ribonucleotide sequence corresponding to a nucleotide sequence of SEQ ID NO: 10; [B] is a ribonucleotide sequence consisting of 3 to 23 nucleotides; and [A'] is a ribonucleotide sequence complementary to [A].